

This listing of claims will replace all prior versions, and listings of claims in the application:

Listing of Claims

1-44. (Canceled)

45. (Original) A method for reducing anxiety in a subject in need thereof by increasing ion flow through KCNQ potassium channels in a cell, the method comprising the step of administering to the subject a pharmaceutical composition comprising a pharmaceutically acceptable carrier and a compound able to increase ion flow through KCNQ potassium channels, said composition administered to the subject in a potassium channel-opening amount, thereby reducing anxiety in the subject.

46. (Original) The method of claim 45, wherein the anxiety is caused by panic disorder, generalized anxiety disorder, or stress disorder.

47. (Original) The method of claim 46, wherein the stress disorder is acute stress disorder or post-traumatic stress disorder.

48. (Original) The method of claim 45, wherein the subject is a human.

49. (Original) The method of claim 45, wherein the KCNQ channel is a heteromeric channel.

50. (Original) The method of claim 45, wherein the KCNQ channel is a homomeric channel.

51. (Original) The method of claim 50, wherein the heteromeric KCNQ channel comprises a KCNQ2 polypeptide subunit.

52. (Original) The method of claim 50, wherein the heteromeric KCNQ channel comprises a KCNQ3 polypeptide subunit.

53. (Original) The method of claim 52, wherein the KCNQ channel is KCNQ2/3.

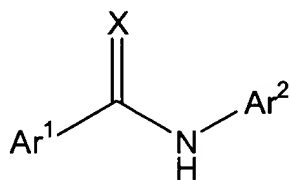
54. (Original) The method of claim 45, wherein the potassium channel-opening amount is 0.1 mg/kg to 200 mg/kg.

55. (Original) The method of claim 54, wherein the potassium channel-opening amount is 10 mg/kg to 100 mg/kg.

56. (Original) The method of claim 45, wherein the composition is administered orally.

57. (Original) The method of claim 45, wherein the composition is administered by injection.

58. (Original) The method of claim 45, wherein the compound able to increase ion flow through KCNQ potassium channels has the formula:



wherein

Ar¹ and Ar² are each members independently selected from the group consisting of aryl, substituted aryl, heteroaryl and substituted heteroaryl; and

X is a member selected from the group consisting of O, S and N-R¹,

wherein R¹ is a member selected from the group consisting of H, (C₁-C₈)alkyl, substituted (C₁-C₈)alkyl, aryl, substituted aryl, heteroaryl, substituted heteroaryl, aryl(C₁-C₄)alkyl, substituted aryl(C₁-C₄)alkyl, CN, -C(O)R², -OR³, -C(O)NR³R⁴, and -S(O)₂NR³R⁴;

wherein R² is a member selected from the group consisting of (C₁-C₈)alkyl, substituted (C₁-C₈)alkyl, aryl, substituted aryl, heteroaryl, substituted heteroaryl, aryl(C₁-C₄)alkyl and substituted aryl(C₁-C₄)alkyl; and R³ and R⁴ are each members independently selected from the group consisting of hydrogen, (C₁-C₈)alkyl, substituted (C₁-C₈)alkyl, aryl, substituted aryl, heteroaryl, substituted heteroaryl, aryl(C₁-C₄)alkyl and substituted aryl(C₁-C₄)alkyl, or R³ and R⁴ can be combined with the nitrogen to which each is attached to form a 5-, 6- or 7-membered ring optionally having additional heteroatoms at the ring vertices.

59. (Original) The method according to claim 58, wherein Ar¹ is a member selected from the group consisting of phenyl, substituted phenyl, indolyl, substituted indolyl, benzofuranyl, substituted benzofuranyl, furanyl, substituted furanyl, thienyl, substituted thienyl, isothiazolyl, substituted isothiazolyl, pyrazolyl and substituted pyrazolyl.

60. (Original) The method according to claim 58, wherein Ar¹ is substituted phenyl, substituted or unsubstituted 2-indolyl and substituted or unsubstituted 2-thienyl.

61. (Original) The method according to claim 58, wherein X is O.

62. (Original) The method according to claim 60, wherein the Ar¹ substituents are selected from the group consisting of halogen, alkyl, halo(C₁-C₄)alkyl, (C₁-C₄)alkoxy, halo(C₁-C₄)alkoxy, nitro, cyano, -NHC(O)R⁷, -NHR⁷, phenyl and substituted phenyl, wherein R⁷ is a member selected from hydrogen, (C₁-C₈)alkyl, substituted (C₁-C₈)alkyl, cycloalkyl, substituted cycloalkyl, heteroalkyl, substituted heteroalkyl, heterocyclyl, substituted heterocyclyl, aryl, substituted aryl, heteroaryl, substituted heteroaryl, aryl(C₁-C₄)alkyl and substituted aryl(C₁-C₄)alkyl, or R⁷ can be combined with the nitrogen to which it is attached to form a 5-, 6- or 7-membered ring optionally having additional heteroatoms at the ring vertices.

63. (Original) The method according to claim 58, wherein Ar² is selected from the group consisting of heteroaryl and substituted heteroaryl.

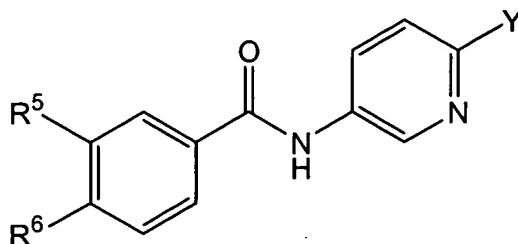
64. (Original) The method according to claim 58, wherein Ar¹ is substituted aryl; Ar² is heteroaryl or substituted heteroaryl; and X is O.

65. (Original) The method according to claim 62, wherein Ar² is pyridyl or substituted pyridyl.

66. (Original) The method according to claim 65, wherein Ar² is selected from the group consisting of 6-methyl-3-pyridyl and 2-chloro-5-pyridyl.

67. (Original) The method according to claim 65, wherein Ar¹ is substituted phenyl.

68. (Original) The method according to claim 67, said compound having the formula:

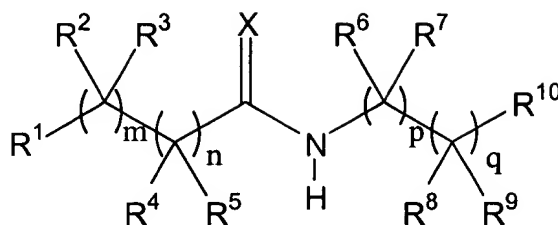


wherein,

Y is a member selected from the group consisting of halogen, C₁-C₄ alkyl, C₁-C₄ substituted alkyl, -OCH₃ and -OCF₃, and R⁵ and R⁶ are members independently selected from the group consisting of H, halogen, alkyl, halo(C₁-C₄)alkyl, nitro, cyano and phenyl, with the proviso that both R⁵ and R⁶ are not H.

69. (Original) The method according to claim 68, wherein R⁵ and R⁶ are members independently selected from the group consisting of H, F, and Cl, with the proviso that both R⁵ and R⁶ are not H.

70. (Original) The method of claim 45, wherein the compound able to increase ion flow through KCNQ potassium channels has the formula:



wherein

R^1 is a member selected from the group consisting of substituted or unsubstituted branched (C_3 - C_8)alkyl, substituted or unsubstituted (C_3 - C_8)cycloalkyl, substituted or unsubstituted (C_3 - C_8)heterocycloalkyl, substituted or unsubstituted aryl and substituted or unsubstituted heteroaryl;

R^2 , R^3 , R^4 and R^5 are each members independently selected from the group consisting of hydrogen, fluorine and substituted or unsubstituted (C_1 - C_8)alkyl, or optionally any two of R^2 , R^3 , R^4 and R^5 are joined together to form a three- to seven-membered ring, having from 0 to 3 heteroatoms as ring members, or R^2 and R^4 taken together form a second bond between the carbon atoms to which each is attached, or R^2 , R^3 , R^4 and R^5 taken together represent a second and third bond between the carbon atoms to which each is attached;

R^6 , R^7 , R^8 and R^9 are each members independently selected from the group consisting of hydrogen, fluorine and substituted or unsubstituted (C_1 - C_8)alkyl, or optionally any two of R^6 , R^7 , R^8 and R^9 are joined together to form a three- to seven-membered ring, having from 0 to 3 heteroatoms as ring members;

R^{10} is a member selected from the group consisting of substituted or unsubstituted (C_3 - C_8)cycloalkyl, substituted or unsubstituted (C_3 - C_8)heterocycloalkyl, substituted or unsubstituted aryl and substituted or unsubstituted heteroaryl;

X is a member selected from the group consisting of O, S and N- R^{11} ,

wherein R^{11} is a member selected from the group consisting of H, (C₁-C₈)alkyl, substituted (C₁-C₈)alkyl, aryl, substituted aryl, heteroaryl, substituted heteroaryl, aryl(C₁-C₄)alkyl, substituted aryl(C₁-C₄)alkyl, -CN, -C(O)R¹², -OR¹³, -NR¹³R¹⁴, -C(O)NR¹³R¹⁴, and -S(O)₂NR¹³R¹⁴;

wherein R^{12} is a member selected from the group consisting of (C₁-C₈)alkyl, substituted (C₁-C₈)alkyl, aryl, substituted aryl, heteroaryl, substituted heteroaryl, aryl(C₁-C₄)alkyl and substituted aryl(C₁-C₄)alkyl; and

R^{13} and R^{14} are each members independently selected from the group consisting of hydrogen, (C₁-C₈)alkyl, substituted (C₁-C₈)alkyl, aryl, substituted aryl, heteroaryl, substituted heteroaryl, aryl(C₁-C₄)alkyl and substituted aryl(C₁-C₄)alkyl, or R^{13} and R^{14} can be combined with the nitrogen to which each is attached to form a 5-, 6- or 7-membered ring optionally having additional heteroatoms at the ring vertices; and

m, n, p and q are each independently an integer of from 0 to 1, with the proviso that at least one of m, n, p or q is 1.

71. (Original) The method of claim 70, wherein X of the compound is O.

72. (Original) The method of claim 70, wherein m and n of the compound are zero.

73. (Original) The method of claim 70, wherein m of the compound is 1 and n of the compound is zero.

74. (Original) The method of claim 70, wherein m and n of the compound are each 1.

75. (Original) The method of claim 70, wherein m and p of the compound are each zero, and n and q of the compound are each 1.

76. (Original) The method of claim 70, wherein m, n, p and q of the compound are each 1.
77. (Original) The method of claim 70, wherein R2 and R4 of the compound, taken together, form a second bond joining the carbon atoms to which each is attached.
78. (Original) The method of claim 70, wherein m and p of the compound are each 1, R2, R3, R6 and R7 of the compound are each hydrogen, n and q of the compound are each zero, and R10 of the compound is selected from the group consisting of substituted or unsubstituted aryl and substituted or unsubstituted heteroaryl.
79. (Original) The method of claim 78, wherein R10 of the compound is substituted aryl having from one to three substituents selected from the group consisting of halogen, halo(C1-C4)alkyl, halo(C1-C4)alkoxy, (C1-C4)alkyl, (C1-C4)alkoxy, nitro, cyano, phenyl and methylenedioxy.
80. (Original) The method of claim 70, wherein m, n, p and q of the compound are each 1, and R2, R3, R4, R5, R6, R7, R8 and R9 of the compound are each hydrogen.
81. (Original) The method of claim 70, wherein m, n, p and q of the compound are each 1; R2, R3, R4, R5, R6, R7, R8 and R9 of the compound are each hydrogen; and R10 of the compound is selected from the group consisting of substituted or unsubstituted aryl and substituted or unsubstituted heteroaryl.
82. (Original) The method of claim 81, wherein R1 of the compound is selected from the group consisting of substituted or unsubstituted branched (C3-C8)alkyl, and substituted or unsubstituted (C3-C8)cycloalkyl.